

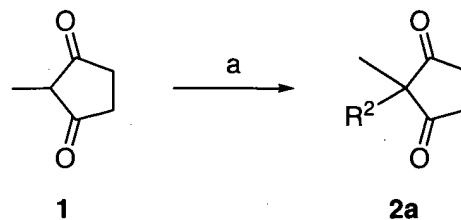


# Replacement Sheet

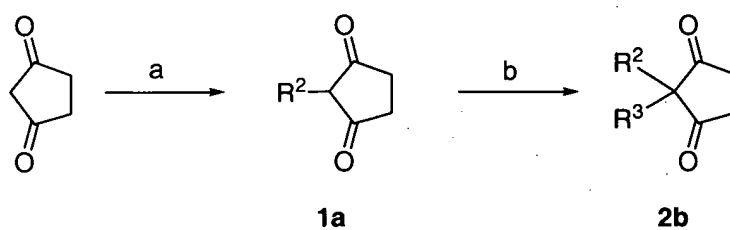
Sheet 1 of 8 DOCKET 17540CIP (AP)  
Serial No. 10/772,720, Applicants: Y. Dondi, et al.  
Title: 10,10-DIALKYL PROSTANOIC ACID DERIVATIVES  
AS AGENTS FOR LOWERING INTRAOCULAR PRESSURE  
714.246.4348 Brent Johnson, ALLERGAN, INC.

Fig. 1

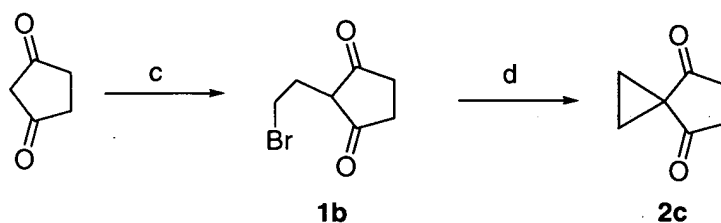
Equation 1



Equation 2



Equation 3

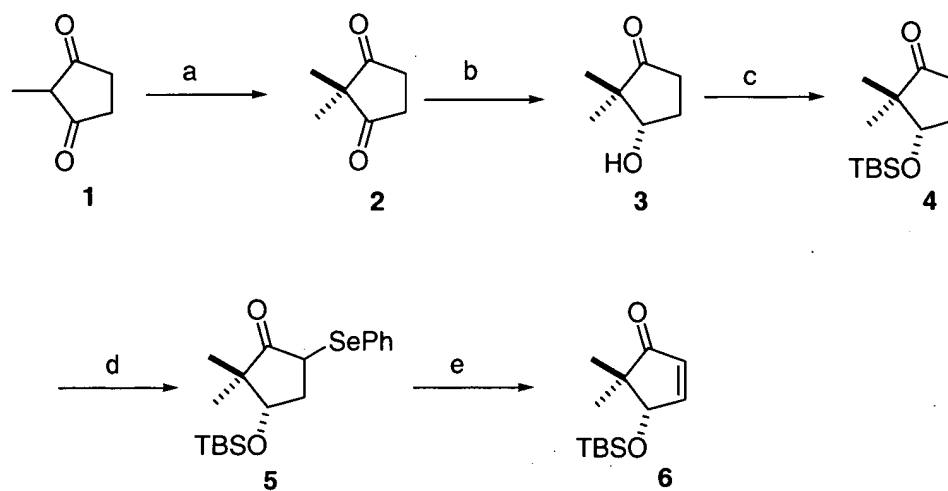


(a) KOH,  $I-R^2$ , dioxane/ $H_2O$ ; (b) KOH,  $I-R^3$ , dioxane/ $H_2O$ ; (c) KOH, 1,2-dibromoethane, dioxane/ $H_2O$ ; (d) KOH, dioxane/ $H_2O$ .

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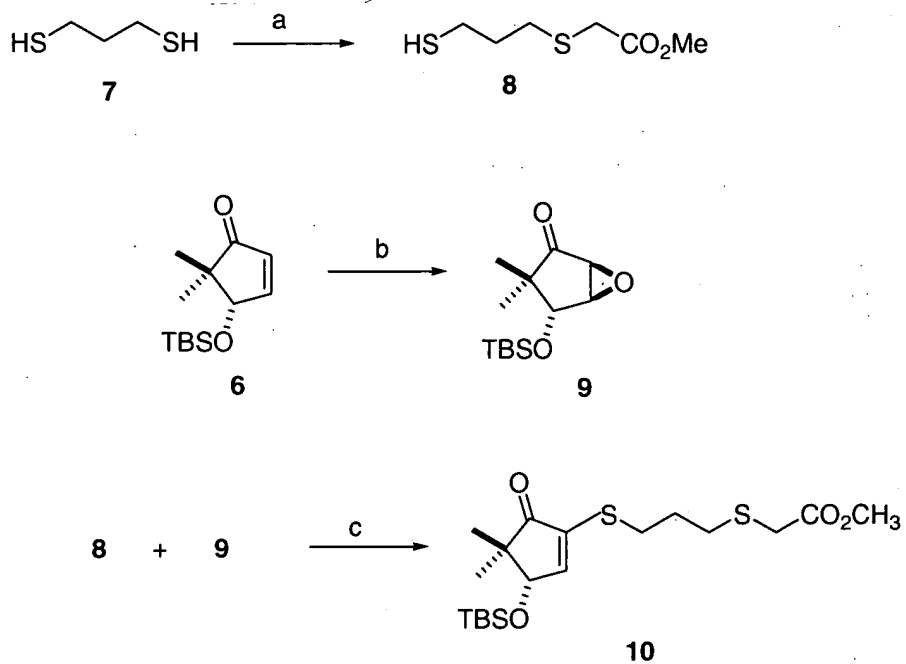
Fig. 2



(a) KOH, MeI, dioxane/H<sub>2</sub>O; (b) Baker's Yeast, D-glucose, H<sub>2</sub>O; (c) TBSOTf, 2,6-lutidine, CH<sub>2</sub>Cl<sub>2</sub>; (d) LDA, THF; PhSeCl; (e) 30% H<sub>2</sub>O<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>.

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Fig. 3

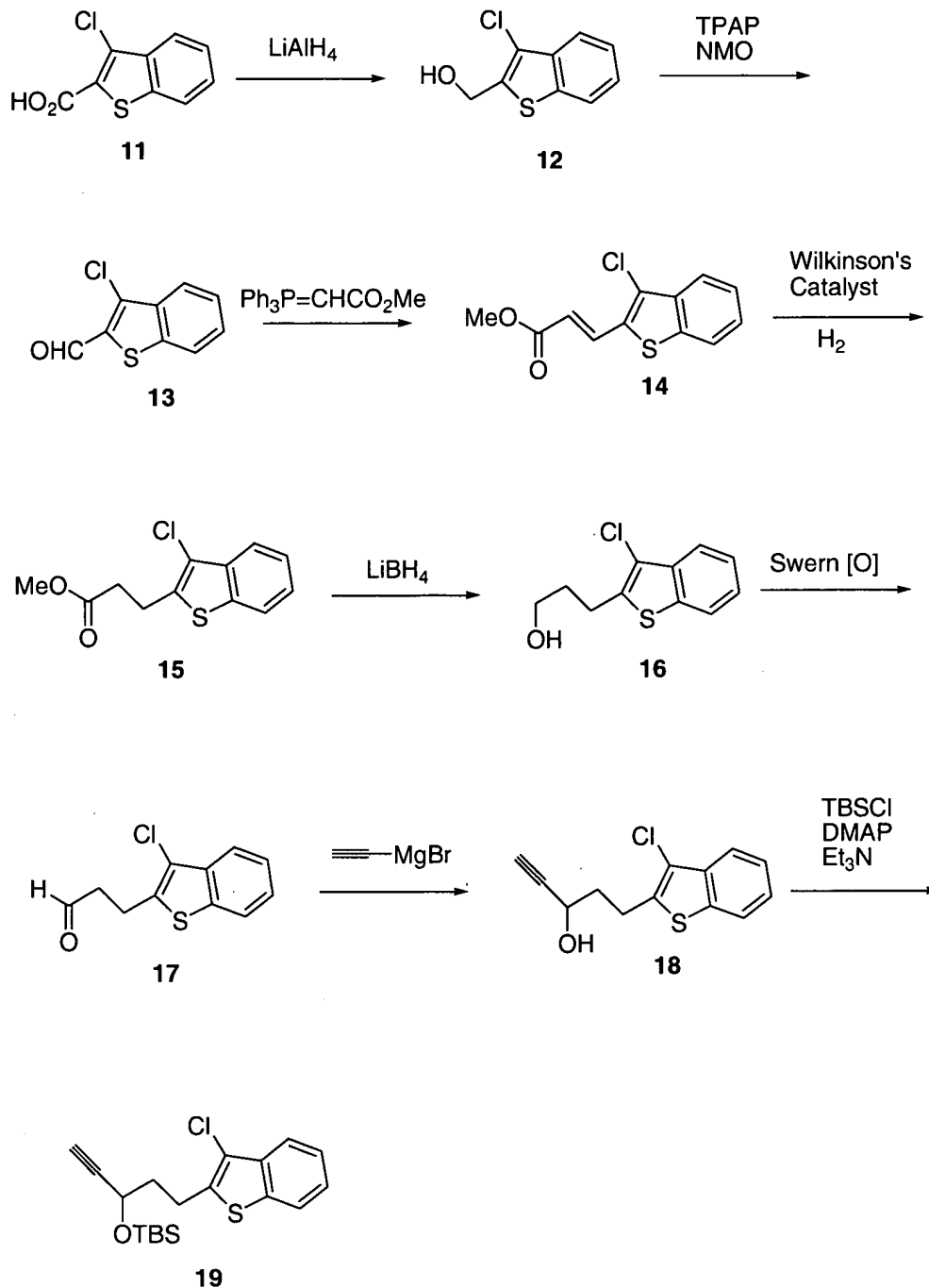


(a) NaH, BrCH<sub>2</sub>CO<sub>2</sub>CH<sub>3</sub>; (b) H<sub>2</sub>O<sub>2</sub>, NaOH, MeOH; (c) basic Alumina, CH<sub>2</sub>Cl<sub>2</sub>.

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Fig. 4



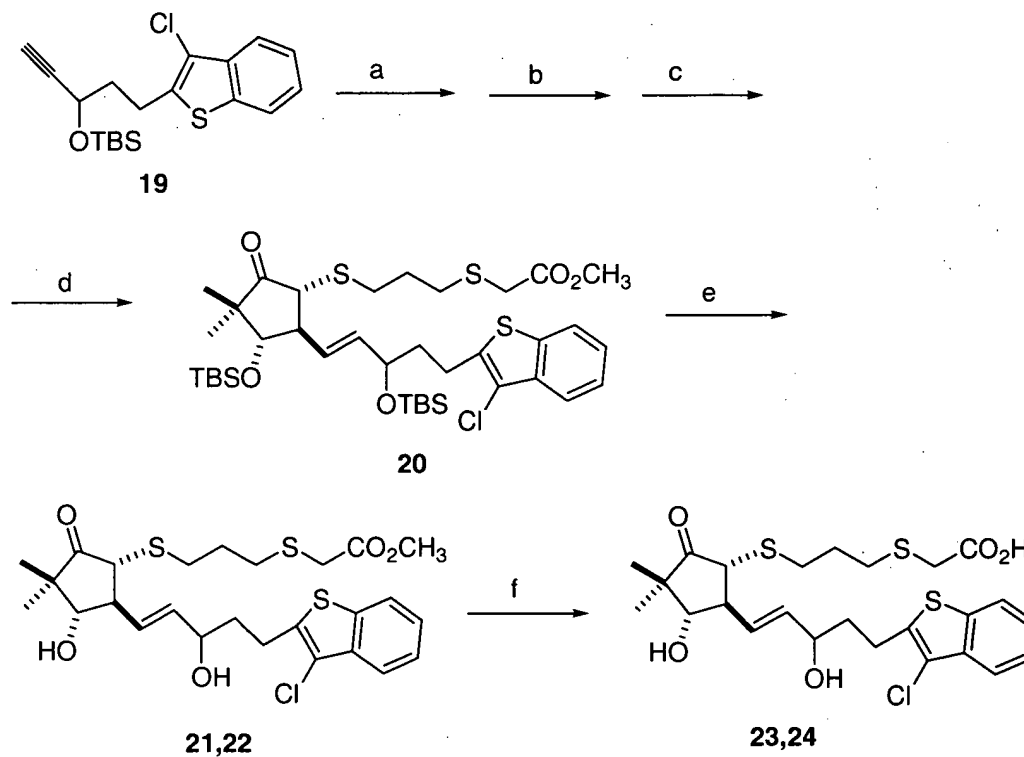
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Fig. 5

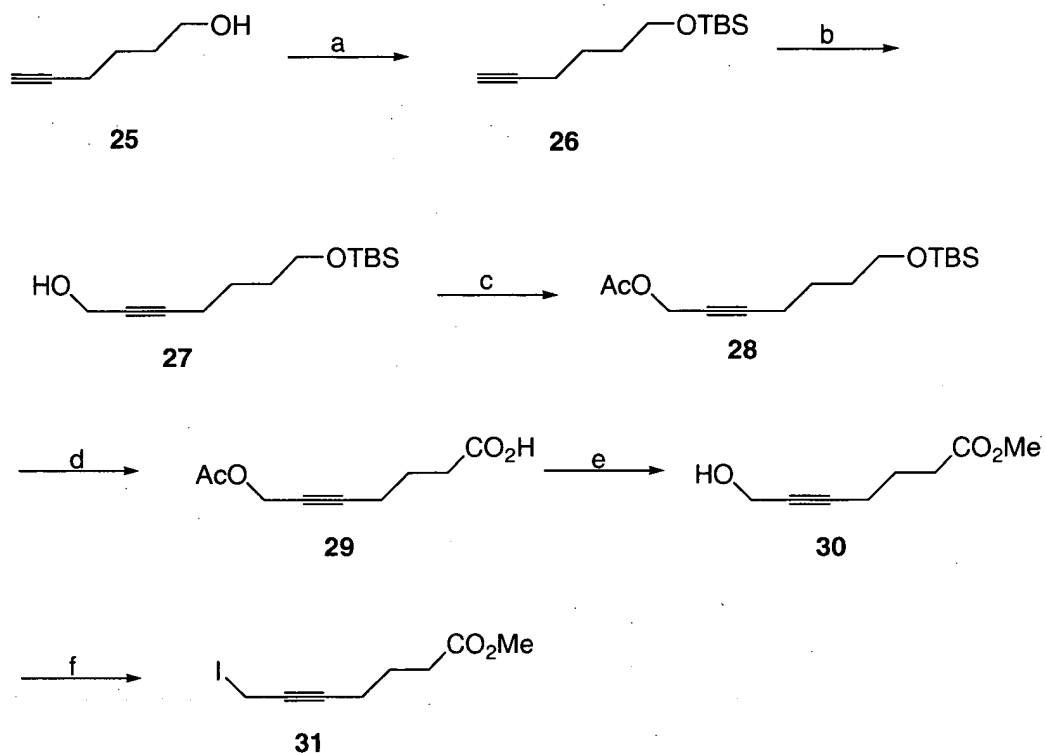


(a)  $\text{Cp}_2\text{ZrHCl}$ , THF; (b) MeLi,  $\text{Et}_2\text{O}$   $-78^\circ\text{C}$ ; (c) lithium 2-thienecyanocuprate;  
(d) enone 10, THF  $-78^\circ\text{C}$ ; (e) HF-pyridine,  $\text{CH}_3\text{CN}$ ; separate diastereomers  
(f) rabbit liver esterase, phosphate buffer,  $\text{CH}_3\text{CN}$ .

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Fig. 6

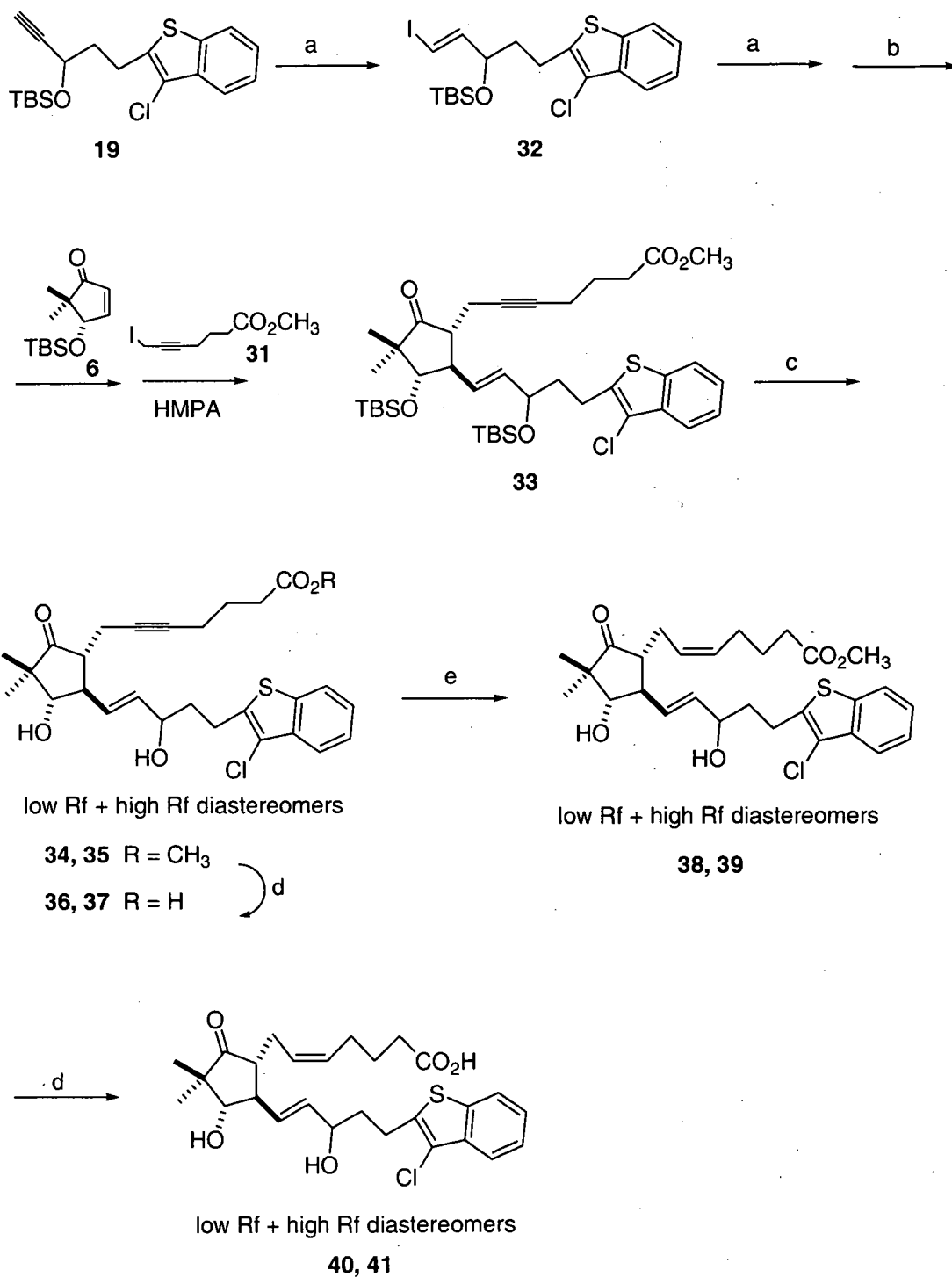


(a) TBSCl, etc.; (b) *n*-BuLi; DMF; (c) Ac<sub>2</sub>O, pyridine; (d) Jones oxidation; (e) MeOH, AcCl; (f) PPh<sub>3</sub>, I<sub>2</sub>, imidazole, CH<sub>2</sub>Cl<sub>2</sub>.

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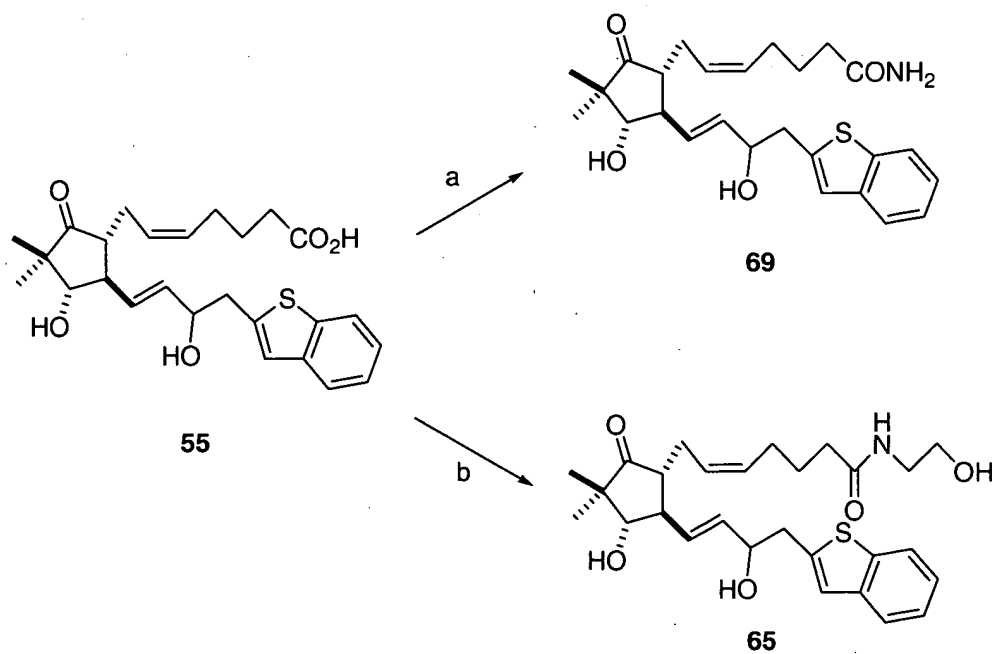
Fig. 7



# Replacement Sheet

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Fig. 8



(a)  $\text{ClCO}_2\text{CH}_2\text{CH}_3$ ,  $\text{Et}_3\text{N}$ ,  $\text{CH}_2\text{Cl}_2$ ;  $\text{NH}_4\text{OH}$  (aq);  
 (b)  $\text{EDCI}$ ,  $\text{N}$ -hydroxysuccinimide,  $\text{H}_2\text{NCH}_2\text{CH}_2\text{OH}$ ,  $\text{DMF}$ .